

10/763,935

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NEWS	12	FEB 25	IMSPRODUCT reloaded with enhancements
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NEWS	14	MAR 31	IFICDB, IFIPAT, and IFIUDB enhanced with new custom IPC display formats
NEWS	15	MAR 31	CAS REGISTRY enhanced with additional experimental spectra
NEWS	16	MAR 31	CA/CAplus and CASREACT patent number format for U.S. applications updated
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NEWS	18	MAR 31	EMBASE, EMBAL, and LEMBASE reloaded with enhancements
NEWS	19	APR 04	STN AnaVist, Version 1, to be discontinued
NEWS	20	APR 15	WPIDS, WPINDEX, and WPIX enhanced with new predefined hit display formats
NEWS	21	APR 28	EMBASE Controlled Term thesaurus enhanced

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L1 19 PP.S.DL..H..RE.L....A.Q.A.QE...R..... /SQSP

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FILE COVERS 1907 - 10 May 2008 VOL 148 ISS 20
FILE LAST UPDATED: 9 May 2008 (20080509/ED)

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=> L1
L2 5 L1

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L2 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2008 ACS on STN
ACCESSION NUMBER: 2007:1480503 HCAPLUS
DOCUMENT NUMBER: 148:254116
TITLE: Peripheral corticotropin releasing factor (CRF) and a novel CRF1 receptor agonist, stressin1-A activate CRF1 receptor expressing cholinergic and nitrergic myenteric neurons selectively in the colon of conscious rats
AUTHOR(S): Yuan, P.-Q.; Million, M.; Wu, S. V.; Rivier, J.; Tache, Y.
CORPORATE SOURCE: CURE: Digestive Diseases Research Center, and Center for Neurovisceral Sciences & Women's Health, VA Greater Los Angeles Healthcare System, Digestive Diseases Division, Department of Medicine and Brain Research Institute, University of California, Los Angeles, Los Angeles, CA, USA
SOURCE: Neurogastroenterology & Motility (2007), 19(11), 923-936
CODEN: NMOTEK; ISSN: 1350-1925
PUBLISHER: Blackwell Publishing Ltd.
DOCUMENT TYPE: Journal
LANGUAGE: English
AB I.p. (i.p.) corticotropin releasing factor (CRF) induced a CRF1 receptor-dependent stimulation of myenteric neurons and motility in the rat proximal colon. We characterize the colonic enteric nervous system response to CRF in conscious rats. Laser capture microdissection combined with reverse transcriptase polymerase chain reaction (RT-PCR) and immunohistochem. in longitudinal muscle myenteric plexus whole-mount colonic prepns. revealed CRF1 receptor expression in myenteric neurons. CRF (i.p., 10 µg kg⁻¹) induced Fos immunoreactivity (IR) (cells per ganglion) selectively in myenteric plexus of proximal (18.3 ± 2.4 vs vehicle: 0.0 ± 0.0) and distal colon (16.8 ± 1.2 vs vehicle: 0.0 ± 0.0), but not in that of gastric corpus, antrum, duodenum, jejunum and ileum. The selective CRF1 agonist, stressin1-A (i.p., 10 µg kg⁻¹)

also induced Fos IR in myenteric but not in submucosal plexus of the proximal and distal colon. Fos IR induced by CRF was located in 55 ± 1.9% and 53 ± 5.1% of CRF1 receptor-IR myenteric neurons and in 44 ± 2.8% and 40 ± 3.9% of cholinergic neurons with Dogiel type I morphol., and in 20 ± 1.6% and 80 ± 3.3% of nitrergic neurons in proximal and distal colon resp. CRF and stressin1-A elicit defecation and diarrhea. These data support that one mechanism through which peripherally injected CRF ligands stimulate colonic function involves a direct action on colonic cholinergic and nitrergic myenteric neurons expressing CRF1 receptor.

IT 935739-46-1, Stressin1 A

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (corticotropin releasing factor1 receptor agonist stressin1-A activated CRF1 expressing cholinergic, nitrergic myenteric neuron in colon eliciting defecation and diarrhea in rat)

RN 935739-46-1 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-L-isoleucyl-, (28→31)-lactam (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

L2 ANSWER 2 OF 5 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2007:1403262 HCPLUS

DOCUMENT NUMBER: 148:70407

TITLE: Subtype-selective corticotropin-releasing factor receptor agonists exert contrasting, but not opposite, effects on anxiety-related behavior in rats

AUTHOR(S): Zhao, Y.; Valdez, G. R.; Fekete, E. M.; Rivier, J. E.; Vale, W. W.; Rice, K. C.; Weiss, F.; Zorrilla, E. P.

CORPORATE SOURCE: Committee on the Neurobiology of Addictive Disorders, The Scripps Research Institute, La Jolla, CA, USA

SOURCE: Journal of Pharmacology and Experimental Therapeutics (2007), 323(3), 846-854

CODEN: JPETAB; ISSN: 0022-3565

PUBLISHER: American Society for Pharmacology and Experimental Therapeutics

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The corticotropin-releasing factor (CRF) system mediates stress responses. Extrahypothalamic CRF1 receptor activation has anxiogenic-like properties, but anxiety-related functions of CRF2 receptors remain unclear. The present study determined the effects of intracerebroventricular administration of a CRF2 agonist, urocortin 3, on behavior of male Wistar rats in the shock-probe, social interaction, and defensive withdrawal tests of anxiety-like behavior. Equimolar doses of stressin1-A, a novel CRF1 agonist, were administered to sep. rats. The effects of pyrazolo[1,5-a]-1,3,5-triazin-4-amine, 8-[4-(bromo)-2-chlorophenyl]-N,N-bis(2-methoxyethyl)-2,7-dimethyl-(9CI) (MJL-1-109-2), a CRF1 antagonist, on behavior in the shock-probe test also were studied. Stressin1-A increased anxiety-like behavior in the social interaction and shock-probe tests. Stressin1-A elicited behavioral activation and defensive burying

at lower doses (0.04 nmol), but it increased freezing, grooming, and mounting at 25-fold higher (1-nmol) doses. Conversely, systemic administration of MJL-1-109-2 (10 mg/kg) had anxiolytic-like effects in the shock-probe test. Unlike stressin1-A or MJL-1-109-2, i.c.v. urocortin 3 infusion did not alter anxiety-like behavior in the shock-probe test across a range of doses that reduced locomotion and rearing and increased grooming. Urocortin 3 also did not decrease social interaction, but it decreased anxiety-like behavior in the defensive withdrawal test at a 2-nmol dose. Thus, i.c.v. administration of CRF1 and CRF2 agonists produced differential, but not opposite, effects on anxiety-like behavior. Urocortin 3 (i.c.v.) did not consistently decrease or increase anxiety-like behavior, the latter unlike effects seen previously after local microinjection of CRF2 agonists into the septum or raphe. With increasing CRF1 activation, however, the behavioral expression of anxiety qual. changes from "coping" to "noncoping" and offensive, agonistic behaviors.

IT 935739-46-1

RL: BSU (Biological study, unclassified); BIOL (Biological study)
(CRF subtype-selective contrasting, but not opposite, effects on anxiety-related behavior in rats)

RN 935739-46-1 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-L-isoleucyl-, (28 \rightarrow 31)-lactam (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 42 THERE ARE 42 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 3 OF 5 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2007:1384929 HCPLUS

DOCUMENT NUMBER: 148:230278

TITLE: Common and Divergent Structural Features of a Series of Corticotropin Releasing Factor-Related Peptides

AUTHOR(S): Grace, Christy Rani R.; Perrin, Marilyn H.; Cantle, Jeffrey P.; Vale, Wylie W.; Rivier, Jean E.; Riek, Roland

CORPORATE SOURCE: Structural Biology Laboratory and The Clayton Foundation Laboratories for Peptide Biology, The Salk Institute for Biological Studies, La Jolla, CA, 92037, USA

SOURCE: Journal of the American Chemical Society (2007), 129(51), 16102-16114

CODEN: JACSAT; ISSN: 0002-7863

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Members of the corticoliberin family include the corticotropin releasing factors (CRFs), sauvagine, the urotensins, and urocortin 1 (Ucn1), which bind to both the CRF receptors CRF-R1 and CRF-R2, and the urocortins 2 (Ucn2) and 3 (Ucn3), which are selective agonists of CRF-R2. Structure activity relationship studies led to several potent and long-acting

analogs with selective binding to either one of the receptors. NMR structures of six ligands of this family (the antagonists astressin B and astressin2-B, the agonists stressin1, and the natural ligands human Ucn1, Ucn2, and Ucn3) were determined in DMSO. These six peptides show differences in binding affinities, receptor-selectivity, and NMR structure. Overall, their backbones are α -helical, with a small kink or a turn around residues 25-27, resulting in a helix-loop-helix motif. The C-terminal helices are of amphipathic nature, whereas the N-terminal helices vary in their amphipathicity. The C-terminal helices thereby assume a conformation very similar to that of astressin bound to the ECD1 of CRF-R2 recently reported by the authors' group. On the basis of an anal. of the observed 3D structures and relative potencies of [Ala]-substituted analogs, it is proposed that both helices could play a crucial role in receptor binding and selectivity. In conclusion, the C-terminal helices may interact along their hydrophobic faces with the ECD1, whereas the entire N-terminal helical surface may be involved in receptor activation. On the basis of the common and divergent features observed in the 3D structures of these ligands, multiple binding models are proposed that may explain their plurality of actions.

IT 1000906-76-2
 RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)
 (common and divergent structural features of series of corticotropin releasing factor-related peptides)

RN 1000906-76-2 HCPLUS
 CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-L-isoleucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-L-isoleucyl-, (28 \rightarrow 31)-lactam (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 67 THERE ARE 67 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 4 OF 5 HCPLUS COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 2007:247482 HCPLUS
 DOCUMENT NUMBER: 146:474752
 TITLE: Stressin1-A, a Potent Corticotropin Releasing Factor Receptor 1 (CRF1)-Selective Peptide Agonist
 AUTHOR(S): Rivier, Jean; Gulyas, Jozsef; Kunitake, Koichi; DiGruccio, Michael; Cantle, Jeffrey P.; Perrin, Marilyn H.; Donaldson, Cindy; Vaughan, Joan; Million, Mulugeta; Gourcerol, Guillaume; Adelson, David W.; Rivier, Catherine; Tache, Yvette; Vale, Wylie
 CORPORATE SOURCE: The Clayton Foundation Laboratories for Peptide Biology, The Salk Institute for Biological Studies, La Jolla, CA, 92037, USA
 SOURCE: Journal of Medicinal Chemistry (2007), 50(7), 1668-1674
 CODEN: JMCMAR; ISSN: 0022-2623
 PUBLISHER: American Chemical Society
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB The potencies and selectivity of peptide CRF antagonists is increased through structural constraints, suggesting that the resulting ligands assume distinct conformations when interacting with CRF1 and CRF2 receptors. To develop selective CRF receptor agonists, we have scanned the sequence -Gln-Ala-His-Ser-Asn-Arg- (residues 30-35 of [DPhe12,Nle21,38]Ac-hCRF4-41) with an i-(i+3) bridge consisting of the Glui-Xaa-Xbb-Lysi+3 scaffold, where residues i = 30, 31, and 32. When i = 31, stressin1-A, a potent CRF1 receptor-selective agonist was generated. In vitro, stressin1-A was equipotent to h/rCRF to release ACTH. Stressin1-A showed a low nanomolar affinity for CRF1 receptor ($K_i = 1.7$ nM) and greater than 100-fold selectivity vs. CRF2 receptor ($K_i = 222$ nM). Stressin1-A released slightly less ACTH than oCRF in adult adrenal-intact male rats, with increased duration of action. Stressin1-A, injected i.p. in rats, induced fecal pellet output (a CRF1 receptor-mediated response) and did not influence gastric emptying and blood pressure (CRF2 receptor-mediated responses).

IT 935739-45-0P 935739-46-1P, Stressin1-A

935739-47-2P 935739-49-4P 935739-51-8P

935739-53-0P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(Stressin1-A as CRF1-selective peptide agonist)

RN 935739-45-0 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-L-isoleucyl- (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 935739-46-1 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-L-isoleucyl-, (28 \rightarrow 31)-lactam (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 935739-47-2 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-N-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-N-methyl-L-leucyl- (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 935739-49-4 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-

L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-N-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-N-methyl-L-leucyl-, (28 \rightarrow 31)-lactam (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 935739-51-8 HCPLUS

CN L-Alaninamide, L-seryl-L-glutaminyl-L- α -glutamyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-threonyl-L-lysyl-L-alanyl-L- α -aspartyl-L-glutaminyl-L-leucyl-L-alanyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-leucyl-L- α -aspartyl-L-isoleucyl-, (31 \rightarrow 34)-lactam (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 935739-53-0 HCPLUS

CN L-Alaninamide, L-seryl-L-glutaminyl-L- α -glutamyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-threonyl-L-lysyl-L-alanyl-L- α -aspartyl-L-glutaminyl-N-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-leucyl-L- α -aspartyl-L-isoleucyl-, (31 \rightarrow 34)-lactam (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

REFERENCE COUNT: 34 THERE ARE 34 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 5 OF 5 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2003:117793 HCPLUS

DOCUMENT NUMBER: 138:153832

TITLE: Preparation of corticotropin-releasing factor (CRF) analogs as CRF receptor type 1 (CRFR1) selective ligands

INVENTOR(S): Rivier, Jean E. F.; Vale, Wylie W., Jr.; Perrin, Marilyn H.; Guylas, Jozsef

PATENT ASSIGNEE(S): The Salk Institute for Biological Studies, USA

SOURCE: PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003011823	A2	20030213	WO 2002-US24238	20020730
WO 2003011823	A3	20070920		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,				

UA, UG, US, UZ, VN, YU, ZA, ZM, ZW
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
 CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
 PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GO, GW, ML, MR,
 NE, SN, TD, TG, AP, EA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, EP, OA
 CA 2455223 A1 20030213 CA 2002-2455223 20020730
 AU 2002355742 A1 20030217 AU 2002-355742 20020730
 JP 2005510458 T 20050421 JP 2003-517015 20020730
 EP 1572679 A2 20050914 EP 2002-752639 20020730
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK
 US 20040204564 A1 20041014 US 2004-763935 20040122
 PRIORITY APPLN. INFO.: US 2001-309504P P 20010801
 WO 2002-US24238 W 20020730

OTHER SOURCE(S): MARPAT 138:153832

AB Corticotropin-releasing factor (CRF) peptides Y1-Pro-Pro-R6-Ser-R8-Asp-R10-R11-D-Phe-R13-R14-R15-Arg-R17-R18-R19-R20-R21-R22-R23-R24-R25-R26-R27-R28-R29-Gln-Glu-R32-R33-R34-Arg-R36-R37-R38-R39-R40-R41-NH₂ (Y1 is acyl having < 15 carbon atoms or radioiodinated tyrosine; the R groups represent various amino acid residues which are defined) or their nontoxic salts are claimed for selective binding to CRFR1. Thus, cyclo(31-34)(Ac-Pro₄,D-Phe₁₂,Nle_{21,38},Glu₃₁,Lys₃₄)-r/hCRF(4-41) was prepared by the solid-phase method and shown to bind hCRFR1 with high affinity and significantly lowered blood pressure when administered peripherally.

IT 496031-18-6P 496031-20-0P 496031-22-2P

496031-24-4P 496031-25-5P

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
 (preparation of corticotropin-releasing factor (CRF) analogs as CRF receptor type 1 (CRFR1) selective ligands)

RN 496031-18-6 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-2-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-2-methyl-L-leucyl- (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 496031-20-0 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-2-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-2-methylalanyl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-2-methyl-L-leucyl- (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 496031-22-2 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-2-

methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-D-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-2-methyl-L-leucyl- (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 496031-24-4 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-2-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-D-alanyl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-2-methyl-L-leucyl- (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 496031-25-5 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-L-isoleucyl- (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

IT 496031-14-2P 496031-15-3P 496031-16-4P
496031-17-5P 496031-19-7P 496031-21-1P
496031-23-3P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of corticotropin-releasing factor (CRF) analogs as CRF receptor type 1 (CRFR1) selective ligands)

RN 496031-14-2 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-L-isoleucyl-, (28 \rightarrow 31)-lactam (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 496031-15-3 HCPLUS

CN L-Leucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-2-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-L-isoleucyl-2-methyl-, (28 \rightarrow 31)-lactam (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 496031-16-4 HCPLUS

CN L-Leucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L-

α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-2-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-2-methylalanyl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-L-isoleucyl-2-methyl-, (28 \rightarrow 31)-lactam (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 496031-17-5 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-2-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-L-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-2-methyl-L-leucyl-, (28 \rightarrow 31)-lactam (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 496031-19-7 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-2-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-2-methylalanyl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-2-methyl-L-leucyl-, (28 \rightarrow 31)-lactam (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 496031-21-1 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-2-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-D-seryl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-2-methyl-L-leucyl-, (28 \rightarrow 31)-lactam (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 496031-23-3 HCPLUS

CN L-Isoleucinamide, 1-acetyl-L-prolyl-L-prolyl-L-isoleucyl-L-seryl-L-leucyl-L- α -aspartyl-L-leucyl-L-threonyl-D-phenylalanyl-L-histidyl-L-leucyl-L-leucyl-L-arginyl-L- α -glutamyl-L-valyl-L-leucyl-L- α -glutamyl-L-norleucyl-L-alanyl-L-arginyl-L-alanyl-L- α -glutamyl-L-glutaminyl-2-methyl-L-leucyl-L-alanyl-L-glutaminyl-L-glutaminyl-L- α -glutamyl-L-histidyl-D-alanyl-L-lysyl-L-arginyl-L-lysyl-L-leucyl-L-norleucyl-L- α -glutamyl-2-methyl-L-leucyl-, (28 \rightarrow 31)-lactam (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

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L1 19 SEA FILE=REGISTRY ABB=ON PLU=ON PP.S.DL..H..RE.L....A.Q.A.QE.
..R...../SQSP

L2 5 SEA FILE=HCPLUS ABB=ON PLU=ON L1

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